

(1) GENERAL INFORMATION:

- (i) APPLICANT: TSYRLOVA, IRENA WOLPE, STEPHEN D.
- (ii) TITLE OF INVENTION: INHIBITOR OF STEM CELL PROLIFERATION AND USES THEREOF
- (iii) NUMBER OF SEQUENCES: 37
- (iv) CORRESPONDENCE ADDRESS:
 - (A) ADDRESSEE: NIXON & VANDERHYE P.C.
 - (B) STREET: 1100 NORTH GLEBE ROAD
 - (C) CITY: ARLINGTON
 - (D) STATE: VIRGINIA
 - (E) COUNTRY: U.S.A.
 - (F) ZIP: 22201-4714
 - (v) COMPUTER READABLE FORM:
 - (A) MEDIUM TYPE: Floppy disk
 - (B) COMPUTER: IBM PC compatible
 - (C) OPERATING SYSTEM: PC-DOS/MS-DOS
 - (D) SOFTWARE: PatentIn Release #1.0, Version #1.30
- (vi) CURRENT APPLICATION DATA:
 - (A) APPLICATION NUMBER: US 08/832,443
 - (B) FILING DATE: 03-APR-1997
 - (C) CLASSIFICATION:
- (viii) ATTORNEY/AGENT INFORMATION:
 - (A) NAME: BYRNE, THOMAS E.
 - (B) REGISTRATION NUMBER: 32,205
 - (C) REFERENCE/DOCKET NUMBER: 1331-222
 - (ix) TELECOMMUNICATION INFORMATION:
 - (A) TELEPHONE: (703) 816-4000
 - (B) TELEFAX: (703) 816-4100
- (2) INFORMATION FOR SEQ ID NO:1:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 13 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear

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(ii) MOLECULE TYPE: peptide

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(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 15 amino acids (B) TYPE: amino acid (C) STRANDEDNESS: (D) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:2: Cys Phe Pro His Phe Asp Leu Ser His Gly Ser Ala Gln Val Cys (2) INFORMATION FOR SEQ ID NO:3: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 19 amino acids (B) TYPE: amino acid (C) STRANDEDNESS: (D) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:3: Asp Ala Leu Thr Asn Ala Val Ala His Val Asp Asp Met Pro Asn Ala 10 Leu Ser Ala (2) INFORMATION FOR SEQ ID NO:4: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 10 amino acids (E) TYPE: amino acid (C) STRANDEDNESS: (D) TOPOLOGY: linear • (ii) MOLECULE TYPE: peptide

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- (C) STRANDEDNESS:
- (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: peptide
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:5:

Leu Val Val Tyr Pro Trp Thr Gln Arg 5

- (2) INFORMATION FOR SEQ ID NO:6:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 8 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: peptide
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:6:

Leu Val Val Tyr Pro Trp Thr Gln 1

- (2) INFORMATION FOR SEQ ID NO:7:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 7 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: peptide
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:7:
 - Leu Val Val Tyr Pro Trp Thr 1
- (2) INFORMATION FOR SEQ ID NO:8:

TI MELECULE TYPE: peptide

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:8: Leu Val Val Tyr Pro Trp (2) INFORMATION FOR SEQ ID NO:9: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 5 amino acids (B) TYPE: amino acid (C) STRANDEDNESS: (D) TOPOLOGY: linear (11) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:9: Leu Val Val Tyr Pro (2) INFORMATION FOR SEQ ID NO:10: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 7 amino acids (B) TYPE: amino acid (C) STRANDEDNESS: (D) TOPOLOGY: linear (1i) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:10: Val Val Tyr Pro Trp Thr Gln 5 (2) INFORMATION FOR SEQ ID NO:11: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 7 amino acids (B) TYPE: amino acid (C) STRANDEDNESS: (D) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide and the second second Typ Fr. Cap The Win Arg Pho

(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 6 amino acids (B) TYPE: amino acid (C) STRANDEDNESS: (D) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:12: Tyr Pro Trp Thr Gln Arg (2) INFORMATION FOR SEQ ID NO:13: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 5 amino acids (B) TYPE: amino acid (C) STRANDEDNESS: (D) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:13: Tyr Pro Trp Thr Gln (2) INFORMATION FOR SEQ ID NO:14: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 6 amino acids (B) TYPE: amino acid (C) STRANDEDNESS: (D) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:14: Ard Met Trp Met Phe Arg

> - Å (LEIDSTH: 413 hase pairs (B) TYPE: nucleic acid

(C) STRANDEDNESS: single

(ii) MOLECULE TYPE: DNA (genomic)

(xi) S	EQUENCE DESC	CRIPTION: SE	EQ ID NO:15:	:		
GTGCTGTCTC	CTGCCGACAA	GACCAACGTC	AAGGCCGCCT	GGGGTAAGGT	CGGCGCGCAC	50
GCTGGCGAGT	ATGGTGCGGA	GGCCCTGGAG	AGGATGTTCC	TGTCCTTCCC	ÇACCACCAAG	120
ACCTACTTCC	CGCACTTCGA	CCTGAGCCAC	GGCTCTGCCC	AGGTTAAGGG	CCACGGCAAG	130
AAGGTGGCCG	ACGCGCTGAC	CAACGCCGTG	GCGCACGTGG	ACGACAT G CC	CAACGCGCTG	240
TCCGCCCTGA	GCGACCTGCA	CGCGCACAAG	CTTCGGGTGG	ACCCGGTCAA	CTTCAAGCTC	300
CTAAGCCACT	GCCTGCTGGT	GACCCTGGCC	GCCCACCTCC	CCGCCGAGTT	CACCCCTGCG	360
GTGCACGCCT	CCCTGGACAA	GTTCCTGGCT	TCTGTGAGCA	CCGTGCTGAC	CTCCAAATAC	420
CGT						423

(2) INFORMATION FOR SEQ ID NO:16:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 141 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: peptide

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:16:

Val Leu Ser Pro Ala Asp Lys Thr Asn Val Lys Ala Ala Trp Gly Lys 1 5 10 15

Tal Gly Ala His Ala Gly Glu Tyr Gly Ala Glu Ala Den Glu Ard Mer 25 30

Phe Leu Ser Phe Pro Thr Thr Lys Thr Tyr Phe Pro His Phe Asp Leu 35 40 45

Ser His Gly Ser Ala Gln Val Lys Gly His Gly Lys Lys Val Ala Asp 50 55 60

Ala Leu Thr Asn Ala Val Ala His Val Asp Asp Met Pro Asn Ala Leu

Ann. Find Type Lee, Lee, Mer Hille Mynches, Lee, Marchine Des Alla Account Lee. (1987)

Leu Ala Ser Val Ser Thr Val Leu Thr Ser Lys Tyr Arg 130 135 140

(2) INFORMATION FOR SEQ ID NO:17:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 438 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: DNA (genomic)
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:17:

GT	GCACCTGA	CTCCTGAGGA	GAAGTCTGCC	GTTACTGCCC	TGTGGGGCAA	GGTGAACGTG	60
GA′	TGAAGTTG	GTGGTGAGGC	CCTGGGCAGG	CTGCTGGTGG	TCTACCTTTG	GACCCAGAGG	120
TT	CTTTGAGT	CCTTTGGGGA	TCTGTCCACT	CCTGATGCTG	TTATGGGCAA	CCCTAAGGTG	180
AA	GGCTCATG	GCAAGAAAGT	GCTCGGTGCC	TTTAGTGATG	GCCTGGCTCA	CCTGGACAAC	240
СТ	CAAGGGCA	CCTTTGCCAC	ACTGAGTGAG	CTGCACTGTG	ACAAGCTGCA	CGTGGATCCT	300
GA(GAACTTCA	GGCTGCTGGG	CAACGTGCTG	GTCTGTGTGC	TGGCCCATCA	CTTTGGCAAA	360
GA	ATTCACCC	CACCAGTGCA	GGCTGCCTAT	CAGAAĀGTGG	TGGCTGGTGT	GGCTAATGCC	420
CTO	GGCCCACA	AGTATCAC					438

- (2) INFORMATION FOR SEQ ID NO:18:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 146 amino acids
 - (B) TYPE: amino acid
 - (€) STRANDEDNESS:
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: peptide
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:18:

Val His Leu Thr Pro Glu Glu Lys Ser Ala Val Thr Ala Leu Trp Gly

.a. Sa. Tyr in Fig. 1m. 4.m Arrine the 4.1 Sec inc. 4.9 Arrine 4

Lys Lys Val Leu Gly Ala Phe Ser Asp Gly Leu Ala His Leu Asp Asn 65 70 75 80

Leu Lys Gly Thr Phe Ala Thr Leu Ser Glu Leu His Cys Asp Lys Leu 85 90 95

His Val Asp Pro Glu Asn Phe Arg Leu Leu Gly Asn Val Leu Val Cys
100 105 110

Val Leu Ala His His Phe Gly Lys Glu Phe Thr Pro Pro Val Gln Ala 115 120 125

Ala Tyr Gln Lys Val Val Ala Gly Val Ala Asn Ala Leu Ala His Lys 130 135 140

Tyr His 145

(2) INFORMATION FOR SEQ ID NO:19:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 141 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: peptide

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:19:

Val Leu Ser Gly Glu Asp Lys Ser Asn Ile Lys Ala Ala Trp Gly Lys 1 5 10 15

Ile Gly Gly His Gly Ala Glu Tyr Gly Ala Glu Ala Leu Glu Arg Met 20 25 30

Phe Ala Ser Phe Pro Thr Thr Lys Thr Tyr Phe Pro His Phe Asp Val 35 40 45

Ser His Gly Ser Ala Gln Val Lys Gly His Gly Lys Lys Val Ala Asp 50 60

Ala Leu Ala Ser Ala Ala Gly His Leu Asp Asp Leu Pro Gly Ala Leu 65 70 75 80

Ser Ala Leu Ser Asp Leu His Ala His Lys Leu Arg Val Asp Pro Val 85 90 95

Test Alla Sem Mail Sea Throttal Lest Thro Sem I va Tyro Aris

(2) INFORMATION FOR SEQ ID NO:20:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 146 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: peptide
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:20:
- Val His Leu Thr Asp Ala Glu Lys Ala Ala Val Ser Cys Leu Trp Gly
 1 5 10 15
- Lys Val Asn Ser Asp Glu Val Gly Glu Ala Leu Gly Arg Leu Leu 20 25 30
- Val Val Tyr Pro Trp Thr Gln Arg Tyr Phe Asp Ser Phe Gly Asp Leu 35 40 45
- Ser Ser Ala Ser Ala Ile Met Gly Asn Ala Lys Val Lys Ala His Gly 50 55 60
- Lys Lys Val Ile Thr Ala Phe Asn Asp Gly Leu Asn His Leu Asp Ser 75 75 80
- Leu Lys Gly Thr Phe Ala Ser Leu Ser Glu Leu His Cys Asp Lys Leu 85 90 95
- His Val Asp Pro Glu Asn Phe Arg Leu Leu Gly Asn Met Ile Val Ile 100 105 110
- Val Leu Gly His His Leu Gly Lys Asp Phe Thr Pro Ala Ala Gln Ala 115 120 125
- Ala Phe Gln Lys Val Val Ala Gly Val Ala Thr Ala Leu Ala His Lys 130 135 140

Tyr His

145

- (2) INFORMATION FOR SEQ ID NO:21:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 141 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:

10 1

Val Gly Gly Gln Ala Gly Ala His Gly Ala Glu Ala Leu Glu Arg Met 20 25 30

Phe Leu Gly Phe Pro Thr Thr Lys Thr Tyr Phe Pro His Phe Asn Leu 35 40 45

Ser His Gly Ser Asp Gln Val Lys Ala His Gly Gln Lys Val Ala Asp 50 55 60

Ala Leu Thr Lys Ala Val Gly His Leu Asp Asp Leu Pro Gly Ala Leu 65 70 75 80

Ser Ala Leu Ser Asp Leu His Ala His Lys Leu Arg Val Asp Pro Val 85 90 95

Asn Phe Lys Leu Leu Ser His Cys Leu Leu Val Thr Leu Ala Ala His
100 105 110

His Pro Asp Asp Phe Asn Pro Ser Val His Ala Ser Leu Asp Lys Phe 115 120 125

Leu Ala Asn Val Ser Thr Val Leu Thr Ser Lys Tyr Arg 130 135 140

(2) INFORMATION FOR SEQ ID NO:22:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 146 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: peptide
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:22:

Val His Leu Ser Ala Glu Glu Lys Glu Ala Val Leu Gly Leu Trp Gly 1 10 15

Lys Val Asn Val Asp Glu Val Gly Glu Ala Leu Gly Arg Leu Leu 20 25 30

Val Val Tyr Pro Trp Thr Gln Arg Phe Phe Glu Ser Phe Gly Asp Leu $35 \hspace{1cm} 40 \hspace{1cm} 45$

Ser Asn Ala Asp Ala Val Met Gly Asn Pro Lys Val Lys Ala His Gly

led Type Bly Inc The Ala Bye Let Dec H. 18. His Typ Asp. And 18.

Val Leu Ala Arg Arg Leu Gly His Asp Phe Asn Pro Asp Val Gln Ala 115 120 Ala Phe Gln Lys Val Val Ala Gly Val Ala Asn Ala Leu Ala His Lys 140 135 Tyr His 145 (2) INFORMATION FOR SEQ ID NO:23: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 23 amino acids (B) TYPE: amino acid (C) STRANDEDNESS: (D) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:23: Val His Leu Ser Ala Glu Glu Lys Glu Ala Val Leu Gly Leu Trp Gly Lys Val Asn Val Asp Glu Val 20 (2) INFORMATION FOR SEQ ID NO:24: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 20 amino acids (B) TYPE: amino acid (C) STRANDEDNESS: (D) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:24: Val Leu Ser Ala Ala Asp Lys Ala Asn Val Lys Ala Ala Trp Gly Lys 10 Val Gly Gly Glr. e ominento a como Botype: amino a mi STRANDEDNESS:
DETORISE VEHICLE DEPOSE

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:25: Phe Pro His Phe Asn Leu Ser His Gly Ser Asp Gln Val Lys 10 (2) INFORMATION FOR SEQ ID NO:26: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 10 amino acids (B) TYPE: amino acid (C) STRANDEDNESS: (T) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:26: Leu Val Val Tyr Pro Trp Thr Gln Arg Phe (2) INFORMATION FOR SEQ ID NO:27: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 8 amino acids (B) TYPE: amino acid (C) STRANDEDNESS: (D) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:27: Val Val Tyr Pro Trp Thr Gln Arg (2) INFORMATION FOR SEQ ID NO:28: (i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 4 amino acids
(B) TYPE: amino acid

Tyr Pro Trp Thr (2) INFORMATION FOR SEQ ID NO:29: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 4 amino acids (B) TYPE: amino acid (C) STRANDEDNESS: (D) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:29: Gly Tyr Pro Tyr (2) INFORMATION FOR SEQ ID NO:30: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 4 amino acids (B) TYPE: amino acid (C) STRANDEDNESS: (D) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:30: Gly Lys Pro Tyr (2) INFORMATION FOR SEQ ID NO:31: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 4 amino acids (B) TYPE: amino acid (C) STRANDEDNESS: (D) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide

(B) TYPE: amino acid (C) STRANDEDNESS: (D) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:32: Lys Met Trp Met Phe Arg (2) INFORMATION FOR SEQ ID NO:33: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 8 amino acids (B) TYPE: amino acid (C) STRANDEDNESS: (D) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:33: Val Arg Arg Met Phe Gly Gly Tyr (2) INFORMATION FOR SEQ ID NO:34: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 6 amino acids (B) TYPE: amino acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear (ii) MONECUNE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:34: Phe Leu Gly Phe Pro Thr 5 (2) INFORMATION FOR SEQ ID NO:35: o organis de la como d ्र । भारत्माताम् सम्बद्धाः १४० र हेर

(ix) FEATURE: (A) NAME/KEY: Modified-site (B) LOCATION: 1 (D) OTHER INFORMATION: /product= "position 1" /label= Xaa /note= "pyroGln" (xi) SEQUENCE DESCRIPTION: SEQ ID NO:35: Xaa Gln Gln Asp Cys Lys (2) INFORMATION FOR SEQ ID NO:36: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 4 amino acids (B) TYPE: amino acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (ix) FEATURE: (A) NAME/KEY: Modified-site (B) LOCATION: 1 (D) OTHER INFORMATION: /product= "position 1" /label= Xaa /note= "N-AcetylSer" (xi) SEQUENCE DESCRIPTION: SEQ ID NO:36: Xaa Asp Lys Pro (2) INFORMATION FOR SEQ ID NO:37: (i) SEQUENCE CHAFACTERISTICS: (A) LENGTH: 13 amino acids (E) TYPE: amino acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (ix) FEATURE: (A) NAME-KEY: Modified-site (E) LOCATION: 1 JA - NAME PETE Modulick bottle

THE OTHER IMPORMANT ME SEE BOTH TERRITOR AT

(E) LOCATION: 4

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:37:

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Xaa Pro His Xaa Asp Leu Ser His Gly Ser Ala Gln Val 1 5 10